Claims:

1. A compound of Formula I, or a salt, solvate, or hydrate thereof

$$R^1$$
 CN
 CN
 R^4

5 wherein

10

15

R¹, R² and R³ are each independently selected from H, OH, C₁₋₆alkyl, C₁₋₆alkoxy, C₁₋₆alkylCO₂, NH₂, NH-C₁₋₆alkyl, N(C₁₋₆alkyl)(C₁₋₆alkyl), C₁₋₆alkyl(C=O)NH, C₁₋₆alkyl(C=O)N(C₁₋₆alkyl), SH, S-C₁₋₆alkyl, NO₂, CF₃, OCF₃ and halo;

R⁴ is unsubstituted Ar, or Ar substituted with 1-4 substituents, independently selected from C₁₋₆alkyl, C₁₋₆alkoxy, and halo;

X is selected from $(CH_2CH_2O)_n$ and $(CH_2)_n$, and n = 1-4.

2. The compound according to claim 1, wherein

R¹, R² and R³ are each independently selected from H, OH, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkylCO₂, NH₂, NH-C₁₋₄alkyl, N(C₁₋₄alkyl)(C₁₋₄alkyl), C₁₋₄alkyl(C=O)NH, C₁₋₄alkyl(C=O)N(C₁₋₄alkyl), NO₂, CF₃, OCF₃, and halo;

 R^4 is C_{1-6} alkyl,

X is $(CH_2CH_2O)_n$, and

n = 1-4.

- 3. The compound according to claim 1 or 2, wherein R¹, R², and R³ are each independently selected from H, OH, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkyl(CO)O, NH₂, NH-C₁₋₄alkyl, N(C₁₋₄alkyl)(C₁₋₄alkyl), C₁₋₄alkyl(C=O)NH, C₁₋₄alkyl(C=O)N(C₁₋₄alkyl), NO₂, CF₃, OCF₃, and halo.
 - 4. The compound according to claim 3, wherein R¹, R² and R³ are each independently selected from H, OH, OCH₃, CH₃CO₂, NH₂, N(CH₃)₂, CH₃CONH, and NO₂.
 - 5. The compound according to claim 4, wherein R¹, R², and R³ are each independently selected from H, OH, and OCH₃.

6. The compound according to claim 1, wherein R⁴ is unsubstituted Ar.

- 7. The compound according to claim 6, wherein R⁴ is phenyl.
- 8. The compound according to claim 2, wherein R⁴ is methyl or ethyl.
- 9. The compound according to claim 8, wherein R^4 is methyl.
- 5 10. The compound according to claim 9, wherein n is 2-3.
 - 11. The compound according to claim 10, wherein n is 3.
 - 12. A compound selected from:
 - 2-Cyano-5-(4-hydroxy-3,5-dimethoxyphenyl)-penta-2E,4E-dienoic acid benzyl ester (CRIX-38)
- 2-Cyano-5-(3,4-dihydroxyphenyl)-penta-2E,4E-dienoic acid benzyl ester (CRIX-39)
 - 2-Cyano-5-(3,4-dihydroxyphenyl)-penta-2E,4E-dienoic acid 2-[2-(2-methoxyethoxy)ethoxy] ethyl ester (CRIV-42)
 - 2-Cyano-5-(4-hydroxy-3,5-dimethoxyphenyl)-penta-2E,4E-dienoic acid 2-[2-(2-methoxyethoxy)ethoxy]ethyl ester (CRIV-46); and
- 2-Cyano-5-(4-hydroxy-3-methoxyphenyl)-penta-2E,4E-dienoic acid benzyl ester (CRIX-79).
 - 13. A composition comprising a compound according to any one of claims 1 to 12 in admixture with a pharmaceutically acceptable diluent or carrier.
- 14. A use of a compound according to any of claims 1-12, and/or a composition according to claim 13, to prepare a medicament to modulate cell proliferation.
 - 15. The use according to claim 14, for inhibiting cell proliferation.
 - 16. The use according to claim 15, wherein the cell is a malignant hematopoietic cell.

17. A method of modulating cell proliferation comprising administering an effective amount of a compound according to any of claims 1-12, and/or a composition according to claim 13, to a cell or animal in need thereof.

- 18. The method according to claim 17, for inhibiting cell proliferation.
- 5 19. The method according to claim 18 wherein the cell is a malignant hematopoietic cell.
 - 20. A compound of Formula III, or a salt, solvate, or hydrate thereof:

wherein

- 10 R¹, R² and R³ are each independently selected from H, OH, C₁₋₆alkyl, C₁₋₆alkoxy, C₁₋₆alkylCO₂, NH₂, NH-C₁₋₆alkyl, N(C₁₋₆alkyl)(C₁₋₆alkyl), C₁₋₆alkyl(C=O)NH, C₁₋₆alkyl(C=O)N(C₁₋₆alkyl), SH, S-C₁₋₆alkyl, NO₂, CF₃, OCF₃, and halo; and R⁴ is selected from C₁₋₆alkyl, phenyl and pyridyl, wherein phenyl and pyridyl are unsubstituted or substituted with 1-4 substituents, independently selected from C₁₋₆alkyl, C₁₋₆alkoxy and halo, with the provisos that when R¹ and R³ are both H and R⁴ is unsubstituted phenyl, R² is not H, Cl, or OCH₃; when R¹ and R² are both H and R⁴ is unsubstituted phenyl, R³ is not NO₂; and when R¹ and R³ are both H and R⁴ is CH₃, R² is not N(CH₃)₂.
- 21. The compound according to claim 1, wherein R¹, R² and R³ are each independently selected from H, OH, C₁₋₄alkyl, C₁₋₄alkoxy, C₁₋₄alkylCO₂, NH₂, NH-C₁₋₄alkyl, N(C₁₋₄alkyl)(C₁₋₄alkyl), C₁₋₄alkyl(C=O)NH, C₁₋₄alkyl(C=O)N(C₁₋₄alkyl), NO₂, CF₃, OCF₃, and halo.
 - 22. The compound according to claim 21, R¹, R² and R³ are each independently selected from H, OH, OCH₃, CH₃CO₂, NH₂, N(CH₃)₂, CH₃CONH, and NO₂.
- 25 23. The compound according to claim 20, wherein R⁴ is selected from C₁₋₄alkyl, phenyl, and pyridyl.

24. The compound according to claim 23, wherein R⁴ is selected from CH₃ and phenyl.

- 25. The compound according to claim 24, wherein R^4 is unsubstituted phenyl.
- 26. The compound according to claim 20, wherein phenyl and pyridyl are unsubstituted or substituted with 1-3 substituents, independently selected from C_{1-4} alkyl, C_{1-4} alkoxy, and halo.
 - 27. The compound according to claim 24, wherein phenyl is unsubstituted or substituted with 1-2 substituents, independently selected from C₁₋₄alkyl, C₁₋₄alkoxy, and halo.
- 10 28. The compound according to claim 20, wherein at least one of \mathbb{R}^1 , \mathbb{R}^2 and \mathbb{R}^3 is OH while \mathbb{R}^4 is selected from unsubstituted phenyl and phenyl substituted with 1-4 substituents, independently selected from C_{1-6} alkyl, C_{1-6} alkoxy, and halo.
 - 29. A compound selected from:
 - 2-Benzenesulfonyl-5-(3,4-dihydroxyphenyl)-penta-2E,4E-dienenitrile (CRVIII-33),
- 2-Benzenesulfonyl-5-(4-hydroxy-3,5-dimethoxyphenyl)-penta-2E,4E-dienenitrile (CRVIII-34),
 - 2-Benzenesulfonyl-5-(4-nitrophenyl)-penta-2E,4E-dienenitrile (CRVIII-35),
 - 5-(4-Acetoxy-3-methoxyphenyl)-2-benzenesulfonyl-penta-2E,4E-dienenitrile (CRVIII-49)
- 5-(3,4-Dihydroxyphenyl)-2-(pyridine-2-sulfonyl)-penta-2E,4E-dienenitrile (CRVIII-50),
 - 2-(4-Chlorobenzenesulfonyl)-5-(3,4-dihydroxyphenyl)-penta-2E,4E-dienenitrile (CRVIII-51),
- 5-(3,4-Dihydroxyphenyl)-2-(toluene-4-sulfonyl)-penta-2E,4E-dienenitrile
 (CRVIII-52), and
 - 5-(3,4-Dihydroxyphenyl)-2-methanesulfonyl-penta-2E,4E-dienenitrile (CRVIII-53).
 - 30. A composition comprising a compound according to any one of claims 20 to 29 in admixture with a pharmaceutically acceptable diluent or carrier.

31. A composition comprising, in admixture with a pharmaceutically acceptable diluent or carrier, a compound of Formula IV, or a salt, solvate, or hydrate thereof

wherein

- R¹, R² and R³ are each independently selected from H, OH, C₁₋₆alkyl, C₁₋₆alkoxy, C₁₋₆alkylCO₂, NH₂, NH-C₁₋₆alkyl, N(C₁₋₆alkyl)(C₁₋₆alkyl), C₁₋₆alkyl(C=O)NH, C₁₋₆alkyl(C=O)N(C₁₋₆alkyl), SH, S-C₁₋₆alkyl, NO₂, CF₃, OCF₃, and halo; and R⁴ is selected from C₁₋₆alkyl, phenyl and pyridyl, wherein phenyl and pyridyl are unsubstituted or substituted with 1-4 substituents, independently selected from C₁₋₆alkyl, C₁₋₆alkoxy, and halo.
 - 32. A use to prepare a medicament to modulate cell proliferation of a composition according to claim 30 or 31, and/or a compound capable of modulating cell proliferation of Formula IV, and/or a salt, solvate or hydrate thereof:

$$R^1$$
 R^2
 CN
 R^3
 IV

15 wherein

20

- R¹, R² and R³ are each independently selected from H, OH, C₁₋₆alkyl, C₁₋₆alkoxy, C₁₋₆alkylCO₂, NH₂, NH-C₁₋₆alkyl, N(C₁₋₆alkyl)(C₁₋₆alkyl), C₁₋₆alkyl(C=O)NH, C₁₋₆alkyl(C=O)N(C₁₋₆alkyl), SH, S-C₁₋₆alkyl, NO₂, CF₃, OCF₃, and halo; and R⁴ is selected from C₁₋₆alkyl, phenyl, and pyridyl, wherein phenyl and pyridyl are unsubstituted or substituted with 1-4 substituents, independently selected from C₁₋₆alkyl, C₁₋₆alkoxy, and halo.
- 33. The use according to claim 13, for inhibiting cell proliferation.
- 34. The use according to claim 14 wherein the cell is a malignant hematopoietic cell.

35. A method of modulating cell proliferation comprising administering to a cell or animal in need thereof an effective amount of a composition according to any of claims 30 and 31, and/or a compound capable of modulating cell proliferation of Formula IV, or a salt, solvate or hydrate thereof:

$$R^1$$
 CN
 R^2
 R^3
 IV

wherein

5

- R¹, R² and R³ are each independently selected from H, OH, C₁₋₆alkyl, C₁₋₆alkoxy, C₁₋₆alkylCO₂, NH₂, NH-C₁₋₆alkyl, N(C₁₋₆alkyl)(C₁₋₆alkyl), C₁₋₆alkyl(C=O)NH, C₁₋₆alkyl(C=O)N(C₁₋₆alkyl), SH, S-C₁₋₆alkyl, NO₂, CF₃, OCF₃, and halo; and R⁴ is selected from C₁₋₆alkyl, phenyl and pyridyl, wherein phenyl and pyridyl are unsubstituted or substituted with 1-4 substituents, independently selected from C₁₋₆alkyl, C₁₋₆alkoxy, and halo.
 - 36. The method according to claim 35, for inhibiting cell proliferation.
- 37. The method according to claim 36, wherein the cell is a malignant hematopoietic cell.